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1	3	(plasminogen adj activator) near2 (IL-2 or IL2 or (IL adj "2"))	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/04/09 13:54
2	56	(IL-2 or IL2 or (IL adj "2")) adj inhibitor	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/04/09 13:54
3	13440	((IL-2 or IL2 or (IL adj "2")) adj inhibitor) near "4" (plasminogen adj activator)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/04/09 13:55
4	1	((IL-2 or IL2 or (IL adj "2")) adj inhibitor) near4 (plasminogen adj activator)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/04/09 13:55
5	1	((IL-2 or IL2 or (IL adj "2")) adj inhibitor) near10 (plasminogen adj activator)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/04/09 13:55

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=> s (plasminogen activator)

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L1 233530 (PLASMINOGEN ACTIVATOR)

=> s (IL-2 or IL2 or (IL 2)) (w) inhibitor

12 FILES SEARCHED...

24 FILES SEARCHED...

36 FILES SEARCHED...

49 FILES SEARCHED...

62 FILES SEARCHED...

67 FILES SEARCHED...

79 FILES SEARCHED...

L2 421 (IL-2 OR IL2 OR (IL 2)) (W) INHIBITOR

=> s l1 (4A) l2

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TI Proteins and nucleic acids encoding same

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PI	US 2004033493	A1	20040219
AI	US 2002-72012	A1	20020131 (10)
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DT Utility
 FS APPLICATION
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 P.C., One Financial Center, Boston, MA, 02111
 CLMN Number of Claims: 49
 ECL Exemplary Claim: 1
 DRWN No Drawings
 LN.CNT 59681
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L5 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2004 ACS on STN DUPLICATE 1
 AN 2001:265240 CAPLUS
 DN 134:261270
 TI **Plasminogen activator** for enhancement of **IL-2 inhibitor** effects
 IN Moriguchi, Akira; Furuichi, Yasuhisa; Katsuta, Kiyotaka; Maeda, Masashi;
 Sato, Natsuki
 PA Fujisawa Pharmaceutical Co., Ltd., Japan
 SO PCT Int. Appl., 21 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2001024784	A2	20010412	WO 2000-JP6874	20001002
	WO 2001024784	A3	20020510		
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PRAI	AU 1999-3249	A	19991004		
	AU 2000-5643	A	20000215		
	WO 2000-JP6874	W	20001002		

AB The present invention is related to a new use of a **plasminogen activator** for increasing an effect caused by **IL-2 inhibitor** and the use of a **IL-2 inhibitor** for increasing or decreasing an effect caused by **plasminogen activator**. Thus, FK506 (Prograf) was administered i.v. by a single bolus injection through the femoral vein 2 h after occlusion of the middle cerebral artery. A tissue-type plasminogen activator (t-PA) (1 mg/kg) was administered i.v. by a bolus injection followed by infusion for 30 min through the femoral vein 2 h after occlusion of the MCA. When drugs were administered 2 h after occlusion of the MCA, FK506 or t-PA showed a relatively small tendency of the inhibition of brain damage. However, the combination of FK506 and t-PA caused the significant reduction of ischemic brain damage and its inhibition is more than 23%, which is greater than that of FK506 or t-PA alone.

=> d 14 1-5 bib ab

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DN 134:261270
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IN Moriguchi, Akira; Furuichi, Yasuhisa; Katsuta, Kiyotaka; Maeda, Masashi; Sato, Natsuki
PA Fujisawa Pharmaceutical Co., Ltd., Japan
SO PCT Int. Appl., 21 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

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	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY				
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PRAI	AU 1999-3249	A	19991004		
	AU 2000-5643	A	20000215		
	WO 2000-JP6874	W	20001002		

AB The present invention is related to a new use of a **plasminogen activator** for increasing an effect caused by **IL-2 inhibitor** and the use of a **IL-2 inhibitor** for increasing or decreasing an effect caused by **plasminogen activator**. Thus, FK506 (Prograf) was administered i.v. by a single bolus injection through the femoral vein 2 h after occlusion of the middle cerebral artery. A tissue-type plasminogen activator (t-PA) (1 mg/kg) was administered i.v. by a bolus injection followed by infusion for 30 min through the femoral vein 2 h after occlusion of the MCA. When drugs were administered 2 h after occlusion of the MCA, FK506 or t-PA showed a relatively small tendency of the inhibition of brain damage. However, the combination of FK506 and t-PA caused the significant reduction of ischemic brain damage and its inhibition is more than 23%, which is greater than that of FK506 or t-PA alone.

L4 ANSWER 2 OF 5 USPATFULL on STN
AN 2004:44501 USPATFULL
TI Proteins and nucleic acids encoding same
IN Tchernev, Velizar T., Branford, CT, UNITED STATES
Spytek, Kimberly A., New Haven, CT, UNITED STATES
Zerhusen, Bryan D., Branford, CT, UNITED STATES
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 Furtak, Katarzyna, Anosia, CT, UNITED STATES
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 Alsobrook, John P., II, Madison, CT, UNITED STATES
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 Rieger, Daniel K., Branford, CT, UNITED STATES
 Burgess, Catherine E., Wethersfield, CT, UNITED STATES

PI	US 2004033493	A1	20040219
AI	US 2002-72012	A1	20020131 (10)
PRAI	US 2001-267459P		20010208 (60)
	US 2001-266975P		20010207 (60)
	US 2001-267057P		20010207 (60)
	US 2001-266767P		20010205 (60)
	US 2001-266406P		20010202 (60)
	US 2001-265395P		20010131 (60)
	US 2001-265412P		20010131 (60)
	US 2001-265517P		20010131 (60)
	US 2001-265514P		20010131 (60)
	US 2001-267823P		20010209 (60)
	US 2001-268974P		20010215 (60)
	US 2001-271855P		20010227 (60)
	US 2001-271839P		20010227 (60)
	US 2001-273046P		20010302 (60)
	US 2001-272788P		20010302 (60)
	US 2001-275989P		20010314 (60)
	US 2001-275925P		20010314 (60)
	US 2001-275947P		20010314 (60)
	US 2001-275950P		20010314 (60)
	US 2001-276450P		20010315 (60)
	US 2001-276448P		20010315 (60)
	US 2001-276397P		20010316 (60)
	US 2001-276768P		20010316 (60)
	US 2001-278652P		20010320 (60)
	US 2001-278775P		20010326 (60)
	US 2001-278778P		20010326 (60)
	US 2001-279882P		20010329 (60)
	US 2001-279884P		20010329 (60)
	US 2001-280147P		20010330 (60)
	US 2001-283083P		20010411 (60)
	US 2001-282992P		20010411 (60)
	US 2001-285133P		20010420 (60)
	US 2001-285749P		20010423 (60)
	US 2001-288327P		20010503 (60)
	US 2001-288504P		20010503 (60)
	US 2001-294047P		20010529 (60)
	US 2001-294473P		20010530 (60)
	US 2001-296964P		20010608 (60)
	US 2001-298959P		20010618 (60)
	US 2001-299324P		20010619 (60)
	US 2001-312020P		20010813 (60)
	US 2001-312908P		20010816 (60)
	US 2001-312889P		20010816 (60)
	US 2001-313930P		20010821 (60)
	US 2001-315470P		20010828 (60)
	US 2001-316447P		20010831 (60)
	US 2001-318115P		20010907 (60)
	US 2001-318118P		20010907 (60)
	US 2001-318740P		20010912 (60)
	US 2001-323379P		20010919 (60)
	US 2001-330308P		20011018 (60)
	US 2001-330245P		20011018 (60)
	US 2001-332701P		20011114 (60)
	US 2001-271664P		20010226 (60)

DT Utility

FS APPLICATION
LREP Ivor R. Elrifi, Ph.D., Mintz, Levin, Cohn, Ferris,, Glovsky and Popeo,
P.C., One Financial Center, Boston, MA, 02111
CLMN Number of Claims: 49
ECL Exemplary Claim: 1
DRWN No Drawings
LN.CNT 59681
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L4 ANSWER 5 OF 5 WPINDEX COPYRIGHT 2004 THOMSON DERWENT on STN

AN 2001-354885 [37] WPINDEX

DNC C2001-109869

TI Use of a plasminogen activator for increasing an effect caused by interleukin-2 inhibitor and for treating acute or chronic cerebral neurodegenerative diseases, e.g. cerebral ischemic diseases and/or brain damage caused by ischemia.

DC B04 D16

IN FURUICHI, Y; KATSUTA, K; MAEDA, M; MORIGUCHI, A; SATO, N

PA (FUJI) FUJISAWA PHARM CO LTD

CYC 21

PI WO 2001024784 A2 20010412 (200137)* EN 21p

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: JP US

EP 1223969 A2 20020724 (200256) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

JP 2003510351 W 20030318 (200321) 25p

ADT WO 2001024784 A2 WO 2000-JP6874 20001002; EP 1223969 A2 EP 2000-963073
20001002, WO 2000-JP6874 20001002; JP 2003510351 W WO 2000-JP6874
20001002, JP 2001-527783 20001002

FDT EP 1223969 A2 Based on WO 2001024784; JP 2003510351 W Based on WO
2001024784

PRAI AU 2000-5643 20000215; AU 1999-3249 19991004

AB WO 200124784 A UPAB: 20010704

NOVELTY - Use of a plasminogen activator for manufacturing a medicament for increasing an effect caused by an IL-2 (interleukin-2) inhibitor, e.g. tacrolimus or its hydrate or a cyclosporin.

DETAILED DESCRIPTION - Use of a plasminogen activator for manufacturing a medicament for increasing an effect caused by an IL-2 (interleukin-2) inhibitor is new.

INDEPENDENT CLAIMS are included for:

(1) a composition comprising a **plasminogen activator** and **IL-2 inhibitor** as a

combined preparation for simultaneous, separate or sequential use for neuroprotective activity;

(2) a manufactured article comprising packaging material containing a plasminogen activator and comprising a label or written material which indicates that the **plasminogen activator** can be used for increasing an effect caused by **IL-2 inhibitor**; and

(3) a manufactured article comprising packaging material containing an IL-2 inhibitor and comprising a label or written material which indicates that the IL-2 inhibitor can be used for increasing or decreasing an effect caused by a plasminogen activator.

ACTIVITY - Neuroprotective; hemostatic; vasotropic; cerebroprotective; cardiatic; thrombolytic; nootropic; anticonvulsant; antiparkinsonian.

Thrombolytic occlusion of the MCA (not defined) was induced in rats and two hours later the **IL-2 inhibitor**

tacrolimus (FK506) and a tissue-type **plasminogen**

activator (t-PA) were administered both alone and in combination.

Alone, FK506 and t-PA showed a relatively small inhibition of brain damage, however the combination of FK506 and t-PA caused significant reduction of ischemic brain damage and produced an inhibition value of 23% (greater than that of FK506 or t-PA alone). Additionally, when the drugs

were administered 3 hours after occlusion of the MCA, t-PA was found to increase the level of brain damage (-13.8 plus or minus 7.0%). In contrast, the combination of FK506 and t-PA caused a significant reduction of ischemic brain damage and produced an inhibition value of 16.2 plus or minus 7.6%. These results show that FK506 is able to decrease the serious damage caused by t-PA.

MECHANISM OF ACTION - The IL-2 inhibitor inhibits the production and activity of IL-2 (claimed). The inhibitor inhibits the transmission of the IL-2 signal.

USE - The **plasminogen activator** and **IL-2 inhibitor** in combination are useful for treating acute or chronic cerebral neurodegenerative diseases, e.g. cerebral ischemic diseases and/or brain damage caused by ischemia (including cerebral infarction, head injury, hemorrhage in the brain such as subarachnoid hemorrhage or intracerebral hemorrhage, cerebral thrombosis, cerebral embolism, cardiac arrest, stroke (such as acute stroke), transient ischemic attacks, hypertensive encephalopathy, Alzheimer's disease, Huntington's disease, Parkinson's disease and amyotrophic lateral sclerosis (ALS). The **IL-2 inhibitor** is useful for increasing or decreasing an effect caused by **plasminogen activator** which comprises a neuroprotective activity or brain damage appeared in case that plasminogen activator is administered after its proper therapeutic time (sic).

ADVANTAGE - Administration of **IL-2 inhibitors** and **plasminogen activators** in combination prolongs the therapeutic time window and also produces increased efficiency and safety in the treatment of ischemic brain damage and administration of an **IL-2 inhibitor** decreases the serious damage caused by a **plasminogen activator**.
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=> d 14 4 bib ab

L4 ANSWER 4 OF 5 USPATFULL on STN
AN 2004:4504 USPATFULL
TI Tumor necrosis factor receptor 2
IN Stanton, Jr., Vincent P., Belmont, MA, United States
PA Nuvelo, Inc., Sunnyvale, CA, United States (U.S. corporation)
PI US 6673908 B1 20040106
AI US 2001-968455 20011001 (9)
RLI Division of Ser. No. US 2000-649035, filed on 25 Aug 2000
Continuation-in-part of Ser. No. US 2000-590749, filed on 8 Jun 2000,
now abandoned Continuation-in-part of Ser. No. US 2000-495780, filed on
1 Feb 2000, now abandoned Continuation-in-part of Ser. No. US
2000-492712, filed on 27 Jan 2000, now abandoned Continuation-in-part of
Ser. No. WO 2000-US1392, filed on 20 Jan 2000 Continuation-in-part of
Ser. No. US 968455 Continuation-in-part of Ser. No. US 1999-451252,
filed on 29 Nov 1999, now abandoned Continuation-in-part of Ser. No. US
1999-427835, filed on 26 Oct 1999, now abandoned Continuation-in-part of
Ser. No. US 1999-414330, filed on 6 Oct 1999, now abandoned
Continuation-in-part of Ser. No. US 1999-389993, filed on 3 Sep 1999,
now abandoned Continuation-in-part of Ser. No. US 1999-370841, filed on
9 Aug 1999, now abandoned Continuation-in-part of Ser. No. US
1999-300747, filed on 26 Apr 1999, now abandoned
PRAI US 1999-131334P 19990426 (60)
US 1999-131191P 19990426 (60)
US 1999-121047P 19990222 (60)
DT Utility
FS GRANTED
EXNAM Primary Examiner: Benzion, Gary; Assistant Examiner: Chakrabarti, Arun
Kr.
LREP Fish & Richardson P.C.
CLMN Number of Claims: 10
ECL Exemplary Claim: 1
DRWN 0 Drawing Figure(s); 0 Drawing Page(s)
LN.CNT 17463
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
AB The present disclosure describes the use of genetic variance information
for genes involved in inflammatory or immunologic disease, disorder, or
dysfunction. The variance information is indicative of the expected
response of a patient to a method of treatment. Methods of determining
relevant variance information and additional methods of using such
variance information are also described.

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